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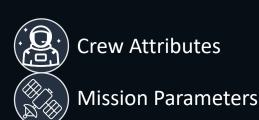
### Motivation

- Demonstrate an approach to incorporating external models with MEDPRAT, enabling us to:
  - Leverage existing work,
  - Improve model fidelity, and
  - Integrate with other risk quantification efforts.



### **MEDPRAT**

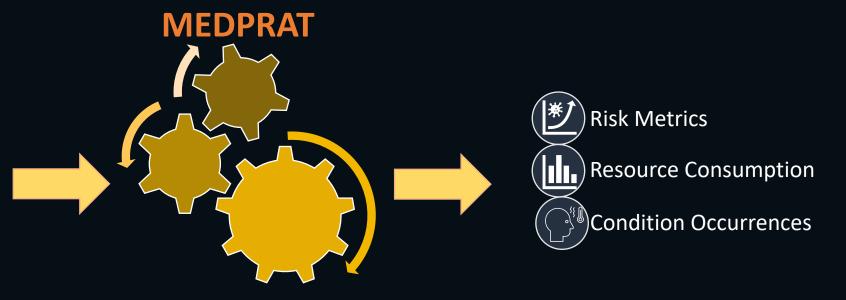
- Medical Extensible Dynamic Probabilistic Risk Assessment Tool
- Computational model that simulates unplanned medical events during a mission to characterize space flight human health and performance risks.







**Treatment Info** 



### Renal Stone Model

 Goodenow-Messman, D.A., Gokoglu, S.A., Kassemi, M. et al. Numerical characterization of astronaut CaOx renal stone incidence rates to quantify in-flight and post-flight relative risk. npj Microgravity 8, 2 (2022).

#### **Urine Chemistry**

 Calcium, Oxalate, Citrate, Magnesium, Uric Acid, Sulfate, Phosphate, Sodium, Potassium, Volume, pH



Biochemical Speciation Model



Renal Stone Size
Distribution



Incidence Rate Ratio

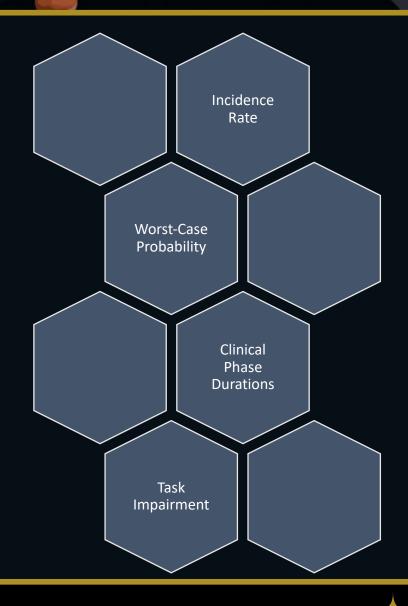




### Integration - Overview

- MEDPRAT offers many "knobs".
  - Flexibility allows for many different types of models to be incorporated.

- Renal Model outputs Incident Rate Ratios (IRRs).
  - Integrate by scaling incidence rate for the nephrolithiasis condition.



### Integration – Challenges With Renal Model

Computational

- Runtime
- Licensing

Data

- Urine Chemistry Details
- Robustness

## Sampling

#### **Input Distributions**

- Select subset of urine chemistry properties
  - Calcium
  - Volume
- Fill in remainder with samples drawn from a similar population



#### IRR Distribution

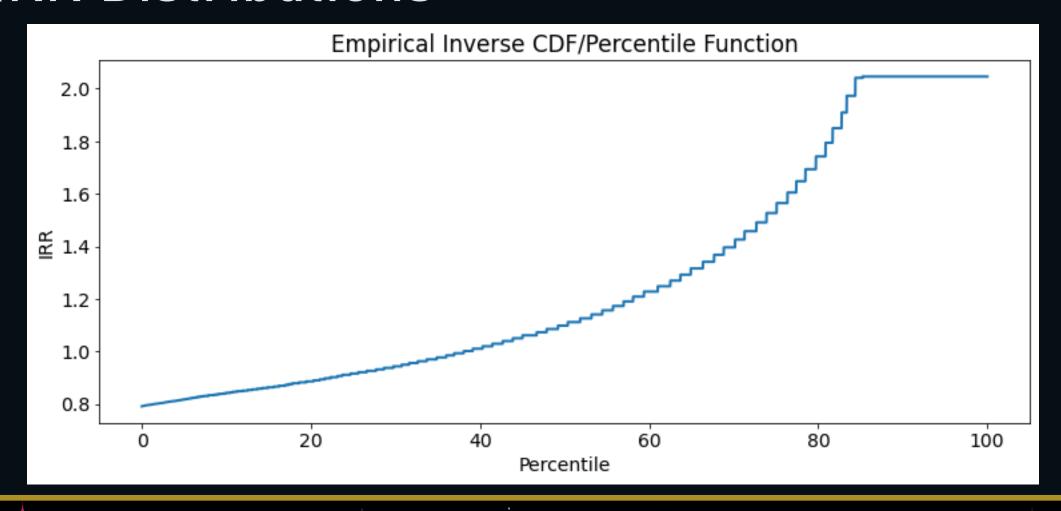
- Measure uncertainty in IRR
- Capture quantities of interest from distribution
  - Mean
  - Median
  - Quantiles



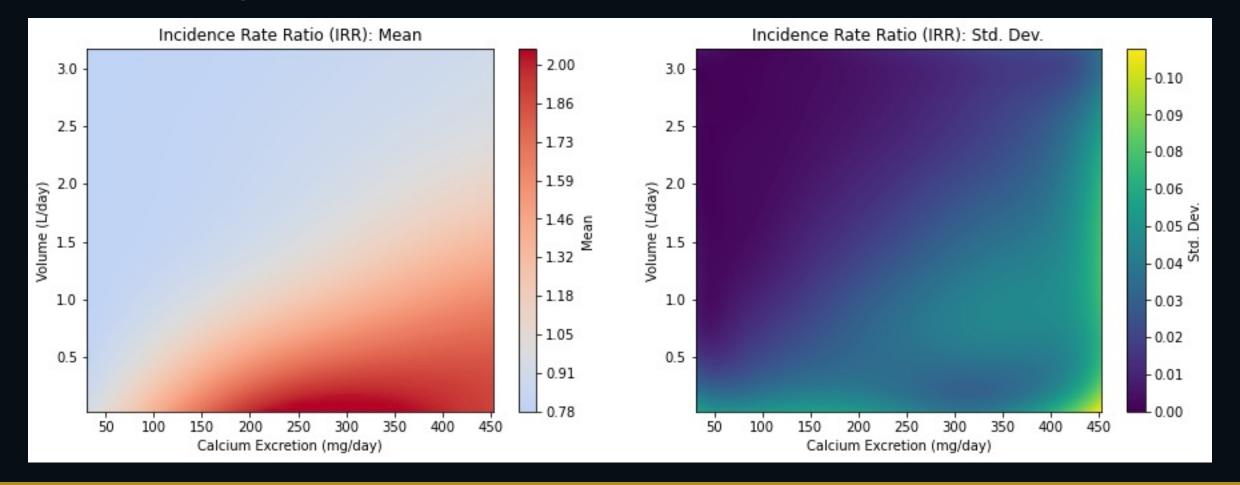




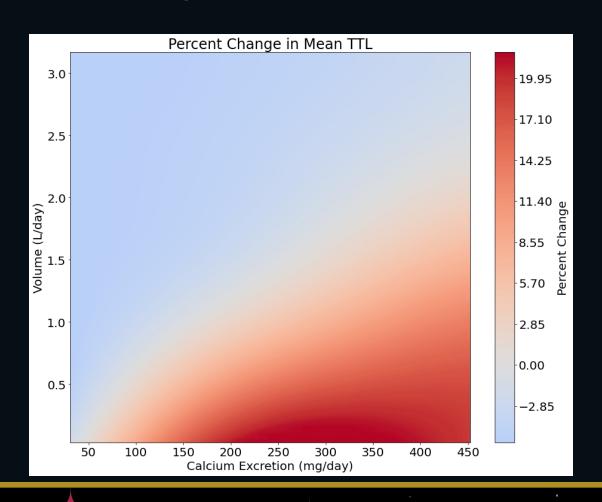
## IRR Distributions

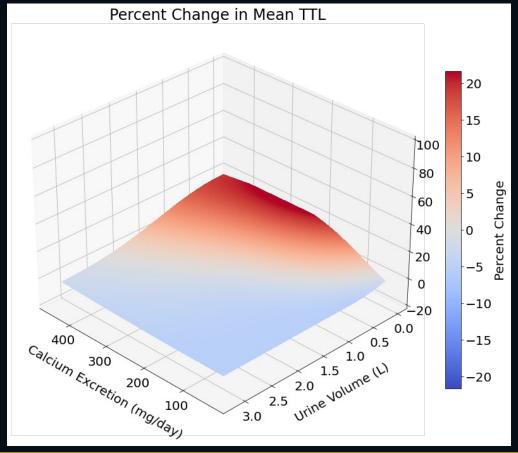


## Surrogate Model

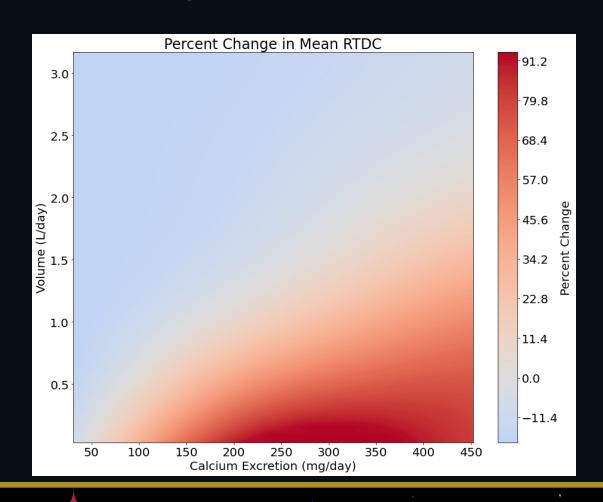


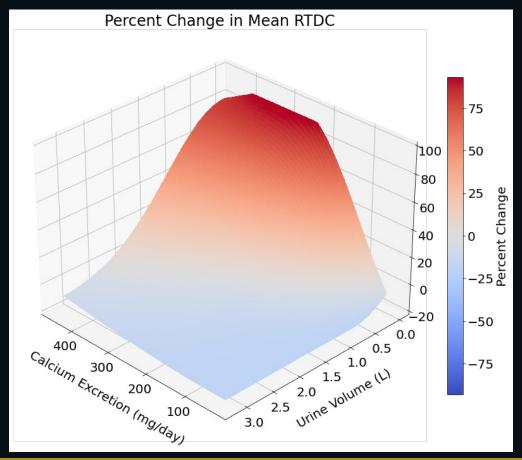
## Change in Risk Metrics - TTL





### Change in Risk Metrics - RTDC







### Conclusions

- Demonstrated a technique for efficiently integrating the results of an external model with MEDPRAT via surrogate modeling of incident rate ratios.
- Incorporated the computed incidence rate changes to evaluate changes in risk metrics (TTL, RTDC).
- This surrogate model is then available for integration with other risk or capability models for holistic risk estimates.





# Thank You

Questions?